

	domains
M	inter-domain sections
-	disulphide bonds
V	variable
C	constant
L	light chain .
Н	heavy chain
Ĺ	constant light chain

FIGURE 2

A.

SEQ ID NO:59 - AME 33 light chain variable region amino acid sequence

EIVLTQSPGTLSLSPGERATLSCRASSSVPYIHWYQQKPGQAPRLLIYATSALASGI PDRFSGSGSGTDFTLTISRLEPEDFAVYYCQQWLSNPPTFGQGTKLEIK

В.

SEQ ID NO:60 - AME 33 light chain variable region nucleic acid sequence

GAAATTGTGTTGACGCAGTCTCCAGGCACCCTGTCTTTGTCTCCAGGGGAAAG
AGCCACCCTCTCCTGCAGGGCCAGCTCAAGTGTACCGTACATCCACTGGTAC
CAGCAGAAACCTGGCCAGGCTCCCAGGCTCCTCATCTATGCCACATCCGCTCT
GGCTTCTGGCATCCCAGACAGGTTCAGTGGCAGTGGGTCTGGGACAGACTTC
ACTCTCACCATCAGCAGACTGGAGCCTGAAGATTTTGCAGTGTATTACTGTCA
GCAGTGGCTGAGTAACCCACCCACTTTTGGCCAGGGGACCAAGCTGGAGATC
AAA

A.

SEQ ID NO:61 - AME 33 heavy chain variable region amino acid sequence

 $\label{thm:condition} EVQLVQSGAEVKKPGESLKISCKGSGRTFTSYNMHWVRQMPGKGLEWMGAIYP\\ LTGDTSYNQKSKLQVTISADKSISTAYLQWSSLKASDTAMYYCARSTYVGGDW\\ QFDVWGKGTTVTVSS$

В.

SEQ ID NO:62 - AME 33 heavy chain variable region nucleic acid sequence

GAGGTGCAGCTGGTGCAGTCTGGAGCAGAGGTGAAAAAGCCCGGGGAGTCT
CTGAAGATCTCCTGTAAGGGTTCTGGCCGTACATTTACCAGTTACAATATGCA
CTGGGTGCGCCAGATGCCCGGGAAAGGCCTGGAGTGGATGGGGGCTATTTAT
CCCTTGACGGGTGATACTTCCTACAATCAGAAGTCGAAACTCCAGGTCACCA
TCTCAGCCGACAAGTCCATCAGCACCGCCTACCTGCAGTGGAGCAGCCTGAA
GGCCTCGGACACCGCCATGTATTACTGTGCGAGATCGACTTACGTGGGCGGT
GACTGGCAGTTCGATGTCTGGGGCAAGGGGACCACGGTCACCGTCTCCTCA

A. Amino acid sequence of a human light chain framework region VkIII (A27) (DPK22) with interspersed CDR sequences labeled

(FRL3 - SEQ ID NO:73) CDRL3 (FRL4 - SEQ ID NO:74)
GIPDRFSGSGSGTDFTLTISRLEPEDFAVYYCXXXXXXXXXX FGQGTKLEIK

B. Nucleic acid sequence of a human light chain framework region VkIII (A27) (DPK22) with interspersed CDR sequences labeled

(FRL1 - SEQ ID NO:75)
GAAATTGTGTTGACGCAGTCTCCAGGCACCCTGTCTTTGTCTCCAGGGAAAGAG

(FRL2 - SEQ ID NO:76) CDRL2
CAGAAACCTGGCCAGGCTCCCAGGCTCCTCATCTAT<u>XXXXXXXXXXXXXXXXXXXX</u>

XXXGGCATCCCAGACAGGTTCAGTGGCAGTGGGTCTGGGACAGACTTCACTCTC

(FRL3 - SEQ ID NO:77) ACCATCAGCAGACTGGAGCCTGAAGATTTTGCAGTGTATTACTGT<u>XXXXXXXXX</u>

A. Amino acid sequence of a human heavy chain framework region VH5-51 (DP-73) with interspersed CDR sequences labeled

(FRH1 - SEQ ID NO:79) CDRH1 (FRH2 - SEQ ID NO:80) EVQLVQSGAEVKKPGESLKISCKGS<u>XXXXXXXXXX</u>WVRQMPGKGLEWMG

CDRH2 (FRH3 - SEQ ID NO:81)

XXXXXXXXXXXXXXXQVTISADKSISTAYLQWSSLKASDTAMYYCAR

CDRH3 (FRH4 - SEQ ID NO:82) XXXXXXXXXXXXWGKGTTVTVSS

B. Nucleic acid sequence of a human heavy chain framework region VH5-51
 (DP-73) with interspersed CDR sequences labeled

(FRHI - SEQ ID NO:83)
GAGGTGCAGCTGGTGCAGTCTGGAGCAGAGGTGAAAAAGCCCGGGGAGTCTCTG

(FRH2- SEQ ID NO:84)
TGGGTGCGCCAGATGCCCGGGAAAGGCCTGGAGTGGATGGGG

CAGGTCACCATCTCAGCCGACAAGTCCATCAGCACCGCCTACCTGCAGTGGAGC

(FRH3 - SEQ ID NO:85) AGCCTGAAGGCCTCGGACACCGCCATGTATTACTGTGCGAGA<u>XXXXXXXXXXX</u>

FIGURE 6

A.

SEQ ID NO:63 - AME 5 light chain variable region amino acid sequence

DIQMTQSPSSLSASVGDRVTITCRASSSVHYIHWYQQKPGKVPKLLIYATSGLAS GVPSRFSGSGSGTDFTLTISSLQPEDVATYYCQTWTFNPPTFGGGTKVEIK

В.

SEQ ID NO:64 - AME 5 light chain variable region nucleic acid sequence

GACATCCAGATGACCCAGTCTCCATCCTCCTGTCTGCATCTGTAGGA
GACAGAGTCACCATCACTTGCAGGGCCAGCTCAAGTGTACATTACATC
CACTGGTACCAGCAGAAACCAGGGAAAGTTCCTAAGCTCTTGATCTAT
GCCACATCCGGCCTGGCTTCTGGGGTCCCATCTCGGTTCAGTGGCAGT
GGATCTGGGACAGATTTCACTCTCACCATCAGCAGCCTGCAGCCTGAA
GATGTTGCCACTTATTACTGCCAGACTTGGACTTTTAACCCTCCCACG
TTCGGCGGAGGGACCAAGGTGGAGATCAAA

A.

SEQ ID NO:65 - AME 5 heavy chain variable region amino acid sequence

 ${\tt QVQLVQSGAEVKKPGASVKVSCKASGYTFTSYNMHWVRQAPGQGLEWMGAIY} \\ {\tt PGNGDTSYNQKFKWRVTMTRDTSTSTVYMELSSLRSEDTAVYYCARSTYYGGD} \\ {\tt WQFDEWGKGTTVTVSS} \\$

В.

SEQ ID NO:66 - AME 5 heavy chain variable region nucleic acid sequence

CAGGTGCAGCTGGTGCAGTCTGGTGCTGAAGTGAAGAAGCCTGGGGCC
TCAGTGAAGGTGTCCTGCAAGGCATCTGGATACACCTTCACCAGCTAC
AATATGCACTGGGTGCGACAGGCCCCTGGACAAGGGCTTGAGTGGATG
GGAGCCATCTATCCTGGAAATGGTGATACAAGCTACAATCAGAAGTTT
AAATGGAGAGTCACCATGACCAGGGACACGTCCACGAGCACAGTCTAC
ATGGAGCTGAGCAGCCTGAGATCTGAGGACACGGCCGTGTATTACTGT
GCGAGATCGACTTATTACGGCGGTGACTGGCAGTTCGACGAGTGGGGC
AAAGGGACCACGGTCACCGTCTCCTCA

FIGURE 8

A. Amino acid sequence of a human light chain framework region VkI (DPK4)

(A20) with interspersed CDR sequences labeled

(FRL1 - SEQ ID NO:87) (CDRL1) (FRL2 - SEQ ID NO:88) (CDRL2) DIQMTQSPSSLSASVGDRVTITC<u>XXXXXXXXXXXXXXX</u>WYQQKPGKVPKLLIY<u>XXXXXXX</u>X

(FRL3 - SEQ ID NO:89) (CDRL3) (FRL4 - SEQ ID NO:90) GVPSRFSGSGSGTDFTLTISSLQPEDVATYYC<u>XXXXXXXXX</u>FGGGTKVEIK

B. Nucleic acid sequence of a human light chain framework region VkI (DPK4) (A20) with interspersed CDR sequences labeled

(FRL1 - SEQ ID NO:91)
GACATCCAGATGACCCAGTCTCCATCCTCCTGTCTGCATCTGTAGGAGACAGAG

(FRL2 - SEQ ID NO:92) (CDRL2)
CAGAAACCAGGGAAAGTTCCTAAGCTCTTGATCTAT<u>XXXXXXXXXXXXXXXXX</u>

(CDRL2 cont.) (FRL3 - SEQ ID NO:93)

XXXXGGGGTCCCATCTCGGTTCAGTGGCAGTGGATCTGGGACAGATTTCACTCT

 ${\tt CACCATCAGCAGCCTGCAGCCTGAAGATGTTGCCACTTATTACTGC} \underline{xxxxxxxx} \\$

TCAAA

A. Amino acid sequence of a human heavy chain framework region VHI DP7/21-2 with interspersed CDR sequences labeled

(FRH1 - SEQ ID NO:95) (CDRH1) (FRH2 - SEQ ID NO:96)
QVQLVQSGAEVKKPGASVKVSCKAS<u>XXXXXXXXXXX</u>WVRQAPGQGLEWMG

(CDRH3) (FRH4 - SEQ ID NO:98) XXXXXXXXXXXXXXWGKGTTVTVSS

B. Nucleic acid sequence of a human heavy chain framework region VHI DP7/21-2 with interspersed CDR sequences labeled

(FRH1 - SEQ ID NO:99)
CAGGTGCAGCTGGTGCAGTCTGGAAGTGAAGAAGCCTGGGGCCTCAGTG

(FRH3 - SEQ ID NO:101)
CAGGGACACGTCCACGAGCACAGTCTACATGGAGCTGAGCAGCCTGAGATCTGA

(CDRH3) (FRH4 - SEQ ID NO:102)

XXXXXXXXXXXXXTGGGGCAAAGGGACCACGGTCACCGTCTCCTCA

FIGURE 10

A.

SEQ ID NO:67 - AME 33 complete light chain amino acid sequence

EIVLTQSPGTLSLSPGERATLSCRASSSVPYIHWYQQKPGQAPRLLIYATSÄLASGIPDR FSGSGSGTDFTLTISRLEPEDFAVYYCQQWLSNPPTFGQGTKLEIK<u>RTVAAPSVFIFPPS</u> DEQLKSGTASVVCLLNNFYPREAKVQWKVDNALQSGNSQESVTEQDSKDSTYSLSST LTLSKADYEKHKVYACEVTHQGLSSPVTKSFNRGEC

- Constant Region is underlined

В.

SEQ ID NO:68 - AME 33 complete light chain nucleic acid sequence

GAAATTGTGTTGACGCAGTCTCCAGGCACCCTGTCTTTGTCTCCAGGGGAAAGAG CCACCCTCTCCTGCAGGGCCAGCTCAAGTGTACCGTACATCCACTGGTACCAGCA GAAACCTGGCCAGGCTCCCAGGCTCCTCATCTATGCCACATCCGCTCTGGCTTCTG GCATCCCAGACAGGTTCAGTGGCAGTGGGTCTGGGACAGACTTCACTCTCACCAT CAGCAGACTGGAGCCTGAAGATTTTGCAGTGTATTACTGTCAGCAGTGGCTGAGT AACCCACCCACTTTTGGCCAGGGGACCAAGCTGGAGATCAAACGAACTGTGGCTG CACCATCTGTCTTCATCTTCCCGCCATCTGATGAGCAGTTGAAATCTGGAACTGCC TCTGTTGTGTGCCTGCTGAATAACTTCTATCCCAGAGAGGCCAAAGTACAGTGGA AGGTGGATAACGCCCTCCAATCGGGTAACTCCCAGGAGAGTGTCACAGAGCAGGA CAGCAAGGACACCTACAGCCTCAGCAGCACCCTGACGCTGAGCAAAGCAGA CTACGAGAAACACAAAGTCTACGCCTGCGAAGTCACCCATCAGGGCCTGAGCTCG CCCGTCACAAAGACTTCAACAGGGGAGAGTGTTAG

FIGURE 11

A. SEQ ID NO:69 - AME 33 complete heavy chain amino acid sequence

EVQLVQSGAEVKKPGESLKISCKGSGRTFTSYNMHWVRQMPGKGLEWMGAIYPLTG
DTSYNQKSKLQVTISADKSISTAYLQWSSLKASDTAMYYCARSTYVGGDWQFDVWG
KGTTVTVSSASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYFPEPVTVSWNSGALTSG
VHTFPAVLQSSGLYSLSSVVTVPSSSLGTQTYICNVNHKPSNTKVDKKVEPKSCDKTH
TCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGV
EVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKA
KGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPPV
LDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK

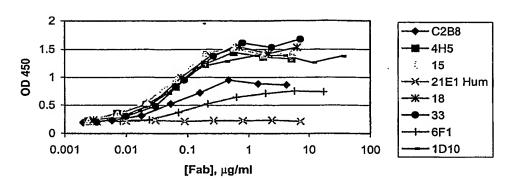
- Constant Region is underlined; the "D" at position 280 and "K" at position 290 are in bold

B. SEQ ID NO:70 - AME 33 complete heavy chain nucleic acid sequence

GAGGTGCAGCTGGAGCAGAGGTGAAAAAGCCCGGGGAGTCTCTG AAGATCTCCTGTAAGGGTTCTGGCCGTACATTTACCAGTTACAATATGCACTGGGT GCGCCAGATGCCCGGGAAAGGCCTGGAGTGGATGGGGGCTATTTATCCCTTGACG GGTGATACTTCCTACAATCAGAAGTCGAAACTCCAGGTCACCATCTCAGCCGACA AGTCCATCAGCACCGCCTACCTGCAGTGGAGCAGCCTGAAGGCCTCGGACACCGC CATGTATTACTGTGCGAGATCGACTTACGTGGGCGGTGACTGGCAGTTCGATGTCT GGGGCAAGGGGACCACGGTCACCGTCTCCTCAGCCTCCACCAAGGGCCCATCGGT CTTCCCCCTGGCACCCTCCCAAGAGCACCTCTGGGGGCACAGCGGCCCTGGGCT GCCTGGTCAAGGACTACTTCCCCGAACCGGTGACGGTGTCGTGGAACTCAGGCGC CCTGACCAGCGCGCGCACACCTTCCCGGCTGTCCTACAGTCCTCAGGACTCTACT CCCTCAGCAGCGTGGTGACCGTGCCCTCCAGCAGCTTGGGCACCCAGACCTACAT CTGCAACGTGAATCACAAGCCCAGCAACACCAAGGTGGACAAGAAGGTTGAGCC CAAATCTTGTGACAAAACTCACACATGCCCACCGTGCCCAGCACCTGAACTCCTG GGGGGACCGTCAGTCTTCCCCCCAAAACCCAAGGACACCCTCATGATCTC CCGGACCCTGAGGTCACATGCGTGGTGGTGGACGTGAGCCACGAAGACCCTGAG GTCAAGTTCAACTGGTACGTGGACGGCGTGGAGGTGCATAATGCCAAGACAAAG CCGCGGGAGGAGCAGTACAACAGCACGTACCGTGTGGTCAGCGTCCTCACCGTCC TGCACCAGGACTGGCTGAATGGCAAGGAGTACAAGTGCAAGGTCTCCAACAAAG CCCTCCCAGCCCCCATCGAGAAACCATCTCCAAAGCCAAAGGGCAGCCCCGAGA ACCACAGGTGTACACCCTGCCCCCATCCCGGGACGAGCTGACCAAGAACCAGGTC AGCCTGACCTGCCTGGTCAAAGGCTTCTATCCCAGCGACATCGCCGTGGAGTGGG AGAGCAATGGGCAGCCGGAGAACAACTACAAGACCACGCCTCCCGTGCTGGACTC CGACGCTCCTTCTTCCTCTATAGCAAGCTCACCGTGGACAAGAGCAGGTGGCAG CAGGGGAACGTCTCCATGCTCCGTGATGCATGAGGCTCTGCACAACCACTACA CGCAGAAGACCTCTCCCTGTCTCCGGGTAAATGA

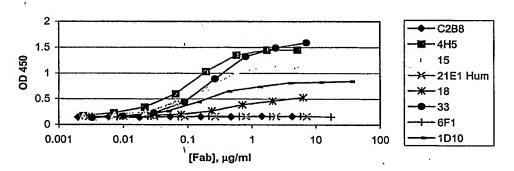
A.

Fab, Binding



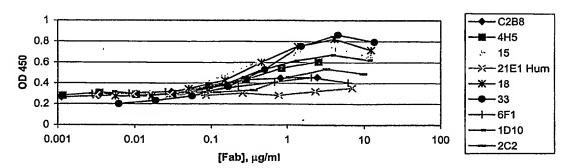
В.

Fab, Off rate



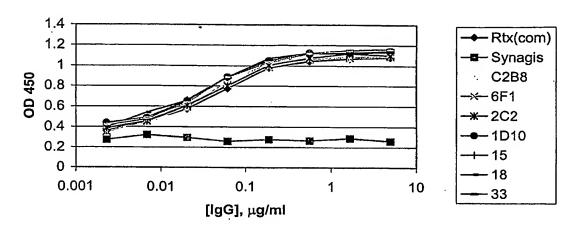
C.

Fab, On rate



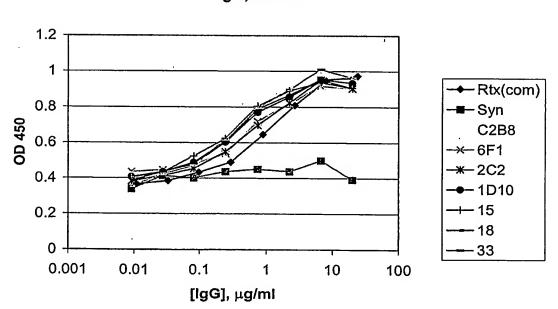
A.

IgG, Off-rate



В.

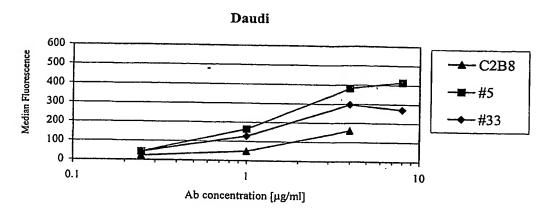
IgG, On-rate



WO 2004/103404

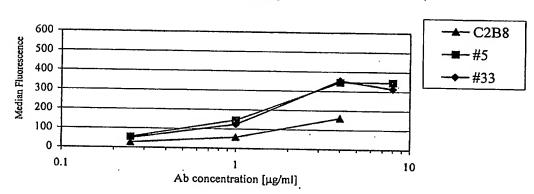
FIGURE 14

A.



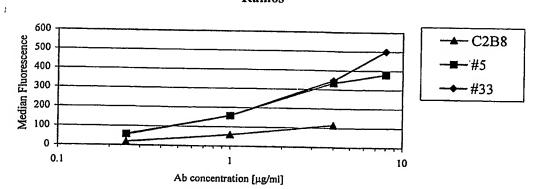
В.

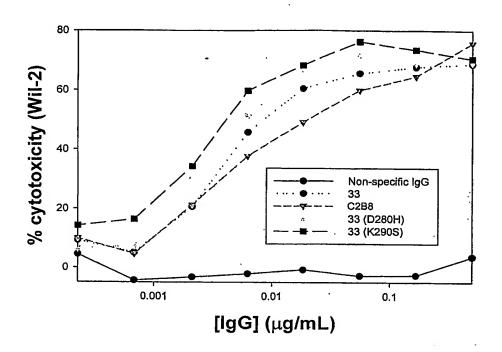
Wil2-S

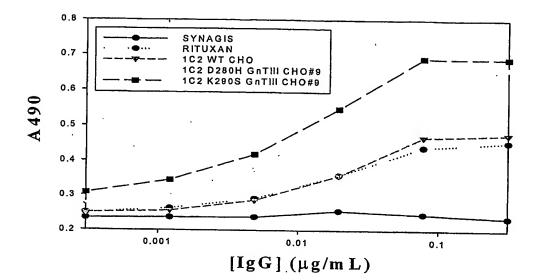


C.

Ramos







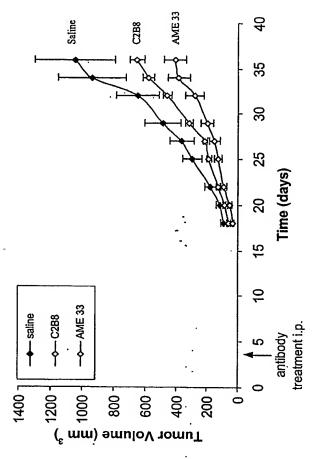


FIGURE 17